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<110> Shi, Yigong <120> CASPASE-9:B

<120> CASPASE-9:BIR3 DOMAIN OF XIAP COMPLEXES AND METHODS OF USE

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<141> 2004-01-30

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Cys Arg Glu Ser Gly Leu Arg Thr Arg Thr Gly Ser Asn Ile Asp Cys 35 40 45

Glu Lys Leu Arg Arg Arg Phe Ser Ser Leu His Phe Met Val Glu Val

Lys Gly Asp Leu Thr Ala Lys Lys Met Val Leu Ala Leu Leu Glu Leu 65 70 75 80

Ala Arg Gln Asp His Gly Ala Leu Asp Cys Cys Val Val Val Ile Leu 85 90 95

Ser His Gly Cys Gln Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr 100 105 110

Gly Thr Asp Gly Cys Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe 115 120 . 125

Asn Gly Thr Ser Cys Pro Ser Leu Gly Gly Lys Pro Lys Leu Phe Phe 130 140

Ile Gln Ala Gys Gly Glu Gln Lys Asp His Gly Phe Glu Val Ala 145 150 155 160 Page 1

Ser Thr Ser Pro Glu Asp Glu Ser Pro Gly Ser Asn Pro Glu Pro Asp 165 170 175

Ala Thr Pro Phe Gln Glu Gly Leu Arg Thr Phe Asp Gln Leu Asp Ala 180 185 190

Ile Ser Ser Leu Pro Thr Pro Ser Asp Ile Phe Val Ser Tyr Ser Thr 195 200 205

Phe Pro Gly Phe Val Ser Trp Arg Asp Pro Lys Ser Gly Ser Trp Tyr 210 215 220

Val Glu Thr Leu Asp Asp Ile Phe Glu Gln Trp Ala His Ser Glu Asp 225 230 235 240

Leu Gln Ser Leu Leu Leu Arg Val Ala Asn Ala Val Ser Val Lys Gly 245 250 255

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Phe His Cys Gly Gly Leu Thr Asp Trp Lys Pro Ser Glu Asp Pro 50 60

Trp Glu Gln His Ala Lys Trp Tyr Pro Gly Cys Lys Tyr Leu Leu Glu 65 70 75 80

Gln Lys Gly Gln Glu Tyr Ile Asn Asn Ile His Leu Thr His Ser Leu 85 90 95

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Gly Ser Arg Arg Asp Gln Ala Arg Gln Leu Ile Ile Asp Leu Glu Thr 50 60

Arg Gly Ser Gln Ala Leu Pro Leu Phe Ile Ser Cys Leu Glu Asp Thr 65 70 75 80

Gly Gln Asp Met Leu Ala Ser Phe Leu Arg Thr Asn Arg Gln Ala Ala 85 90 95

Lys Leu Ser Lys Pro Thr Leu Glu Asn Leu Thr Pro Val Val Leu Arg 100 105 110

Pro Glu Ile Arg Lys Pro Glu Val Leu Arg Pro Glu Thr Pro Arg Pro 115 120 125

Val Asp Ile Gly Ser Gly Gly Phe Gly Asp Val Gly Ala Leu Glu Ser 130 140

Leu Arg Gly Asn Ala Asp Leu Ala Tyr Ile Leu Ser Met Glu Pro Cys 145 150 155 160

Gly His Cys Leu Ile Ile Asn Asn Val Asn Phe Cys Arg Glu Ser Gly 165 170 175

Leu Arg Thr Arg Thr Gly Ser Asn Ile Asp Cys Glu Lys Leu Arg Arg 180 185 190

Arg Phe Ser Ser Leu His Phe Met Val Glu Val Lys Gly Asp Leu Thr 195 200 205 Page 3

Ala Lys Lys Met Val Leu Ala Leu Leu Glu Leu Ala Gln Gln Asp His 210

Gly Ala Leu Asp Cys Cys Val Val Val Ile Leu Ser His Gly Cys Gln 220

Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr Gly Thr Asp Gly Cys 245 250 255

Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe Asn Gly Thr Ser Cys 260 265 270

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Ser Trp Arg Asp Pro Lys Ser Gly Ser Trp Tyr Val Glu Thr Leu Asp 355 360 365

Asp Ile Phe Glu Gln Trp Ala His Ser Glu Asp Leu Gln Ser Leu Leu 370 380

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Gly Gln Asp Met Leu Ala Ser Phe Leu Arg Thr Asn Arg Gln Ala Ala 85 90 95

Lys Leu Ser Lys Pro Thr Leu Glu Asn Leu Thr Pro Val Val Leu Arg 100 105 110

Pro Glu Ile Arg Lys Pro Glu Val Leu Arg Pro Glu Thr Pro Arg Pro 115 120 125 Page 5

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Gly Ser Arg Arg Asp Gln Ala Arg Gln Leu Ile Ile Asp Leu Glu Thr 50 60

Arg Gly Ser Gln Ala Leu Pro Leu Phe Ile Ser Cys Leu Glu Asp Thr 65 70 75 80

Gly Gln Asp Met Leu Ala Ser Phe Leu Arg Thr Asn Arg Gln Ala Ala 85 90 95

Lys Leu Ser Lys Pro Thr Leu Glu Asn Leu Thr Pro Val Val Leu Arg 100 105 110

Pro Glu Ile Arg Lys Pro Glu Val Leu Arg Pro Glu Thr Pro Arg Pro 115 120 125

Val Asp Ile Gly Ser Gly Gly Phe Gly Asp Val Gly Ala Leu Glu Ser 130 135 140

Leu Arg Gly Asn Ala Asp Leu Ala Tyr Ile Leu Ser Met Glu Pro Cys 145 150 155 160

Gly His Cys Leu Ile Ile Asn Asn Val Asn Phe Cys Arg Glu Ser Gly 165 170 175

Leu Arg Thr Arg Thr Gly Ser Asn Ile Asp Cys Glu Lys Leu Arg Arg Page 7

Arg Phe Ser Ser Leu His Phe Met Val Glu Val Lys Gly Asp Leu Thr 195 200 205 Ala Lys Lys Met Val Leu Ala Leu Leu Glu Leu Ala Gln Gln Asp His 210 220 Gly Ala Leu Asp Cys Cys Val Val Val Ile Leu Ser His Gly Cys Gln 225 230 235 240 Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr Gly Thr Asp Gly Cys 245 250 255 Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe Asn Gly Thr Ser Cys 260 265 270 Ser Leu Gly Gly Lys Pro Lys Leu Phe Phe Ile Gln Ala Cys Gly 275 280 285 Gly Glu Gln Lys Asp His Gly Phe Glu Val Ala Ser Thr Ser Pro Glu 290 295 300 Asp Glu Ser Pro Gly Ser Asm Pro Glu Pro Asp Ala Thr Pro Phe Gln 305 310 315 320 Glu Gly Leu Arg Thr Phe Asp Gln Leu Asp Ala Ile Ser Ser Leu Pro 325 330 335 Thr Pro Ser Asp Ile Phe Val Ser Tyr Ser Thr Phe Pro Gly Phe Val 340 345 350 Ser Trp Arg Asp Pro Lys Ser Gly Ser Trp Tyr Val Glu Thr Leu Asp 355 360 365 Asp Ile Phe Glu Gln Trp Ala His Ser Glu Asp Leu Gln Ser Leu Leu 370 380 Leu Arg Val Ala Asn Ala Val Ser Val Lys Gly Ile Tyr Lys Gln Met 385 390 395 400 Pro Gly Cys Phe Asn Phe Leu Arg Lys Leu Phe Phe Lys Thr Ser 405 410 415

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Ile Ile Asn Asn Lys Asn Phe His Lys Ser Thr Gly Met Thr Ser Arg 50 55 60

Ser Gly Thr Asp Val Asp Ala Ala Asn Leu Arg Glu Thr Phe Arg Asn 65 70 75 80

Leu Lys Tyr Glu Val Arg Asn Lys Asn Asp Leu Thr Arg Glu Glu Ile 85 90 95

Val Glu Leu Met Arg Asp Val Ser Lys Glu Asp His Ser Lys Arg Ser 100 105 110

Phe Val Cys Val Leu Leu Ser His Gly Glu Glu Gly Ile Ile Phe 115 120 125

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Gln Ala Cys Arg Gly Thr Glu Leu Asp Cys Gly Ile Glu Thr Asp Ser 165 170 175

Gly Val Asp Asp Asp Met Ala Cys His Lys Ile Pro Val Glu Ala Asp 180 185 190

Phe Leu Tyr Ala Tyr Ser Thr Ala Pro Gly Tyr Tyr Ser Trp Arg Asn

Ser Lys Asp Gly Ser Trp Phe Ile Gln Ser Leu Cys Ala Met Leu Lys 210 220

Gln Tyr Ala Asp Lys Leu Glu Phe Met His Ile Leu Thr Arg Val Asn 235 230 240

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Val Arg Asn Lys Asn Asp Leu Thr Arg Glu Glu Ile Val Glu Leu Met 65 70 75 80

Arg Asp Val Ser Lys Glu Asp His Ser Lys Arg Ser Ser Phe Val Cys 85 90 95

Val Leu Leu Ser His Gly Glu Glu Gly Ile Ile Phe Gly Thr Asn Gly $100 \hspace{1cm} 105 \hspace{1cm} 110$

Pro Val Asp Leu Lys Lys Ile Thr Asn Phe Phe Arg Gly Asp Arg Cys 115 120 125

Arg Ser Leu Thr Gly Lys Pro Lys Leu Phe Ile Ile Gln Ala Cys Arg Page 10 Gly Thr Glu Leu Asp Cys Gly Ile Glu Thr Asp Ser Gly Val Asp Asp 145 150 155 160

135

Asp Met Ala Cys His Lys Ile Pro Val Glu Ala Asp Phe Leu Tyr Ala 165 170 175

Tyr Ser Thr Ala Pro Gly Tyr Tyr Ser Trp Arg Asn Ser Lys Asp Gly 180 185 190

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Lys Leu Glu Phe Met His Ile Leu Thr Arg Val Asn Arg Lys Val Ala 210 215 220

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Leu Gly Phe Asp Val Ile Val Tyr Asn Asp Cys Ser Cys Ala Lys Met Page 11 Gln Asp Leu Lys Lys Ala Ser Glu Glu Asp His Thr Asn Ala Ala 100 105 110

Cys Phe Ala Cys Ile Leu Leu Ser His Gly Glu Glu Asn Val Ile Tyr 115 120 125

Gly Lys Asp Gly Val Thr Pro Ile Lys Asp Leu Thr Ala His Phe Arg 130 135 140

Gly Asp Arg Cys Lys Thr Leu Leu Glu Lys Pro Lys Leu Phe Phe Ile 145 150 155 160

Gln Ala Cys Arg Gly Thr Glu Leu Asp Asp Gly Ile Gln Ala Asp Ser 165 170 175

Gly Pro Ile Asn Asp Thr Asp Ala Asn Pro Arg Tyr Lys Ile Pro Val 180 185 190

Glu Ala Asp Phe Leu Phe Ala Tyr Ser Thr Val Pro Gly Tyr Tyr Ser 195 200 205

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Ile Leu Glu Glu His Gly Lys Asp Leu Glu Ile Met Gln Ile Leu Thr 225 230 235 240

Arg Val Asn Asp Arg Val Ala Arg His Phe Glu Ser Gln Ser Asp Asp 245 250 255

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420 425 430 Lys Glu Ile Ser Thr Glu Glu Gln Leu Arg Arg Leu Gln Glu Glu Lys 435 440 445 Leu Cys Lys Ile Cys Met Asp Arg Asn Ile Ala Ile Val Phe Val Pro 450 460 Cys Gly His Leu Val Thr Cys Lys Gln Cys Ala Glu Ala Val Asp Lys 465 470 475 480 Cys Pro Met Cys Tyr Thr Val Ile Thr Phe Lys Gln Lys Ile Phe Met 485 490 495

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Arg Met Ser Thr Tyr Ser Thr Phe Pro Ala Gly Val Pro Val Ser Glu 50 60

Arg Ser Leu Ala Arg Ala Gly Phe Tyr Tyr Thr Gly Val Asn Asp Lys 65 70 75 80

Val Lys Cys Phe Cys Cys Gly Leu Met Leu Asp Asn Trp Lys Leu Gly 85 90 95

Asp Ser Pro Ile Gln Lys His Lys Gln Leu Tyr Pro Ser Cys Ser Phe $100 \hspace{1cm} 105 \hspace{1cm} 110$

Ile Gln Asn Leu Val Ser Ala Ser Leu Gly Ser Thr Ser Lys Asn Thr 115 120 125

Ser Pro Met Arg Asn Ser Phe Ala His Ser Leu Ser Pro Thr Leu Glu 130 140

His Ser Ser Leu Phe Ser Gly Ser Tyr Ser Ser Leu Ser Pro Asn Pro 145 150 155 160

Leu Asn Ser Arg Ala Val Glu Asp Ile Ser Ser Ser Arg Thr Asn Pro 165 170 . 175

Tyr Ser Tyr Ala Met Ser Thr Glu Glu Ala Arg Phe Leu Thr Tyr His 180 185 190

Met Trp Pro Leu Thr Phe Leu Ser Pro Ser Glu Leu Ala Arg Ala Gly 195 200 205

Phe Tyr Tyr Ile Gly Pro Gly Asp Arg Val Ala Cys Phe Ala Cys Gly 210 215 220

Gly Lys Leu Ser Asn Trp Glu Pro Lys Asp Asp Ala Met Ser Glu His 225 230 235 240

Arg Arg His Phe Pro Asn Cys Pro Phe Leu Glu Asn Ser Leu Glu Thr 245 Leu Arg Phe Ser Ile Ser Asn Leu Ser Met Gln Thr His Ala Ala Arg 260 265 270 Met Arg Thr Phe Met Tyr Trp Pro Ser Ser Val Pro Val Gln Pro Glu Gln Leu Ala Ser Ala Gly Phe Tyr Tyr Val Gly Arg Asn Asp Asp Val 290 295 300 Lys Cys Phe Cys Cys Asp Gly Gly Leu Arg Cys Trp Glu Ser Gly Asp 305 310 315 320 Asp Pro Trp Val Glu His Ala Lys Trp Phe Pro Arg Cys Glu Phe Leu 325 330 335 Ile Arg Met Lys Gly Gln Glu Phe Val Asp Glu Ile Gln Gly Arg Tyr 340 345 350 Pro His Leu Leu Glu Gln Leu Leu Ser Thr Ser Asp Thr Thr Gly Glu 355 360 365 Glu Asn Ala Asp Pro Pro Ile Ile His Phe Gly Pro Gly Glu Ser Ser 370 375 380 Ser Glu Asp Ala Val Met Met Asn Thr Pro Val Val Lys Ser Ala Leu 385 390 395 400 Glu Met Gly Phe Asn Arg Asp Leu Val Lys Gln Thr Val Gln Ser Lys 405 410 415 Ile Leu Thr Thr Gly Glu Asn Tyr Lys Thr Val Asn Asp Ile Val Ser 420 425 430 Ala Leu Leu Asn Ala Glu Asp Glu Lys Arg Glu Glu Glu Lys 435 Gln Ala Glu Glu Met Ala Ser Asp Asp Leu Ser Leu Ile Arg Lys Asn 450 455 460 Arg Met Ala Leu Phe Gln Gln Leu Thr Cys Val Leu Pro Ile Leu Asp 465 470 475 480 Asn Leu Leu Lys Ala Asn Val Ile Asn Lys Gln Glu His Asp Ile Ile 485 490 495 Page 16

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Ile Leu Val Lys Gly Asn Ala Ala Ala Asn Ile Phe Lys Asn Cys Leu 515 520 525

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Gln Leu Arg Arg Leu Gln Glu Glu Arg Thr Cys Lys Val Cys Met Asp 565 570 575

Lys Glu Val Ser Val Val Phe Ile Pro Cys Gly His Leu Val Val Cys 580 585 590

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Met Ser Thr Tyr Ser Thr Phe Pro Ala Gly Val Pro Val Ser Glu Arg 35 40 45

Ser Leu Ala Arg Ala Gly Phe Tyr Tyr Thr Gly Val Asn Asp Lys Val 50 55 60

Lys Cys Phe Cys Cys Gly Leu Met Leu Asp Asn Trp Lys Arg Gly Asp 65 70 75 80

Ser Pro Thr Glu Lys His Lys Lys Leu Tyr Pro Ser Cys Arg Phe Val 85 90 95 Page 17

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Asp His Val Asp Gly Gln Ile Leu Gly Gln Leu Arg Pro Leu Thr Glu 50 60

Glu Glu Glu Glu Gly Ala Gly Ala Thr Leu Ser Arg Gly Pro Ala 65 70 75 80

Phe Pro Gly Met Gly Ser Glu Glu Leu Arg Leu Ala Ser Phe Tyr Asp 85 90 95

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Asp Phe Val His Ser Val Gln Glu Thr His Ser Gln Leu Leu Gly Ser 165 170 175

Trp Asp Pro Trp Glu Glu Pro Glu Asp Ala Ala Pro Val Ala Pro Ser 180 185 190

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Ala Glu Ala Ala Tyr Gln Thr Gly Ala Asp Gln Ala Ser Ile Thr Ala 115 120 125

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Glu Leu Phe Arg Pro His Met Ile Glu Asp Ile Gln Arg Ala Gly Ser Page 24

Gly Ser Arg Arg Asp Gln Ala Arg Gln Leu Ile Ile Asp Leu Glu Thr $50 \hspace{1cm} 55 \hspace{1cm} 60$ Arg Gly Ser Gln Ala Leu Pro Leu Phe Ile Ser Cys Leu Glu Asp Thr 65 70 75 80 Gly Gln Asp Met Leu Ala Ser Phe Leu Arg Thr Asn Arg Gln Ala Ala 85 90 95 Lys Leu Ser Lys Pro Thr Leu Glu Asn Leu Thr Pro Val Val Leu Arg 100 105 110 Pro Glu Ile Arg Lys Pro Glu Val Leu Arg Pro Glu Thr Pro Arg Pro 115 120 125 Val Asp Ile Gly Ser Gly Gly Phe Gly Asp Val Gly Ala Leu Glu Ser 130 140 Leu Arg Gly Asn Ala Asp Leu Ala Tyr Ile Leu Ser Met Glu Pro Cys 145 150 155 160 Gly His Cys Leu Ile Ile Asn Asn Val Asn Phe Cys Arg Glu Ser Gly 165 170 175 Leu Arg Thr Arg Thr Gly Ser Asn Ile Asp Cys Glu Lys Leu Arg Arg 180 185 190 Arg Phe Ser Ser Leu His Phe Met Val Glu Val Lys Gly Asp Leu Thr 195 200 205 Ala Lys Lys Met Val Leu Ala Leu Leu Glu Leu Ala Gln Gln Asp His 210 220 Gly Ala Leu Asp Cys Cys Val Val Ile Leu Ser His Gly Cys Gln 225 230 235 240 Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr Gly Thr Asp Gly Cys 245 250 255 Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe Asn Gly Thr Ser Cys 260 265 270 Pro Ser Leu Gly Gly Lys Pro Lys Leu Phe Phe Ile Gln Ala Cys Gly 275 280 285

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Gly Thr Asp Gly Cys Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe 115 120 125

Asn Gly Thr Ser Cys Pro Ser Leu Gly Gly Lys Pro Lys Leu Phe Phe 130 140

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Ser His Gly Cys Gln Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr 100 105 110

Gly Thr Asp Gly Cys Pro Val Ser Val Glu Lys Ile Val Asn Ile Phe 115 120 125

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Glu Lys Leu Arg Arg Arg Phe Ser Ser Leu His Phe Met Val Glu Val 50 60
Lys Gly Asp Leu Thr Ala Lys Lys Met Val Leu Ala Leu Leu Glu Leu 65 70 75 80
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Ser His Gly Cys Gln Ala Ser His Leu Gln Phe Pro Gly Ala Val Tyr
100 105 110
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Phe	Phe	Lys 275	Thr	Ser											



CERTIFICATE OF MAILING UNDER 37 C.F.R. § 1.8

In re Application of:

Shi

Serial No. 10/769,218

Group Art Unit: 1646

Filed:

January 30, 2004

Examiner: not yet assigned

For:

CASPACE-9:BIR3 DOMAIN OF XIAP COMPLEXES AND METHODS

OF USE

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JENNIFER MARTINEZ

(Typed or printed name of person mailing paper or fee)

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5. Certificate of Mailing